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(21) International Application Number: PCT/US00/05879 (22) International Filing Date: 7 March 2000 (07.03.00) (30) Priority Data: 60/124,306 12 March 1999 (12.03.99) US 60/158,201 7 October 1999 (07.10.99) US (71) Applicant: AMERICAN CYANAMID COMPANY [US/US]; Five Giralda Farms, Madison, NJ 07940 (US). (72) Inventors: TREACY, Michael, Frank; 53 Sequoia Drive, Newtown, PA 18940 (US). BORYSEWICZ, Raymond, Frank; 12 Albemarle Road, Hamilton Square, NJ 08690 (US). SCHWINGHAMMER, Kurt, Allen; 1206 Uni- versity Drive, Yardley, PA 19067 (US). RENSNER, Paul, Erich; 1267 Woodthrush Court, Yardley, PA 19067 (US). OLOUMI-SADEGHI, Hassan; 1204 Goldenrod Court, Yardley, PA 19067 (US). (74) Agents: HOGAN, John, W.; American Home Products Corpo- ration, Patent Law Dept. 2B2, One Campus Drive, Parsip- pany, NJ 07054 (US) et al.		(81) Designated States: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG). Published <i>Without international search report and to be republished upon receipt of that report.</i>
(54) Title: SYNERGISTIC INSECTICIDAL COMPOSITIONS (57) Abstract The present invention provides a synergistic insecticidal composition comprising as essential active ingredients a neuronal sodium channel antagonist in combination with one or more compounds selected from the group consisting of pyrethroids, pyrethroid-type compounds, recombinant nucleopolyhedroviruses capable of expressing an insect toxin, organophosphates, carbamates, formamidines, macrocyclic lactones, amidinohydrazones, GABA antagonists and acetylcholine receptor ligands. Also provided are methods for synergistic insect control and crop protection.		

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SYNERGISTIC INSECTICIDAL COMPOSITIONS

BACKGROUND OF THE INVENTION

Insecticidal agents and compositions have been developed to control insect pests such as agrohorticultural pests, hygienic pests, or wood-eating pests and in practice have been used as a single or a
5 mixed agent. However, economically efficient and ecologically safe insect control compositions are still being sought. Insecticidal compositions which allow for reduced effective dosage rates, increased environmental safety and lower incidence of insect
10 resistance are highly desirable. Although the rotational application of insect control agents having different modes of action may be adopted for good pest management practice, this approach does not necessarily give satisfactory insect control. Further, even though
15 combinations of insect control agents have been studied, a high synergistic action has not always been found. Obtaining an insecticidal composition which demonstrates no cross-resistance to existing insecticidal agents, no toxicity problems and little
20 negative impact on the environment is extremely difficult.

Therefore, it is an object of this invention to provide a synergistic insecticidal composition which demonstrates a high controlling effect with concomittant
25 reduced crop production cost and reduced environmental load.

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It is another object of this invention to provide methods for synergistic insect control and enhanced crop protection.

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SUMMARY OF THE INVENTION

The present invention provides a synergistic insecticidal composition comprising as essential active ingredients a synergistically effective amount of a neuronal sodium channel antagonist in combination with one or more compounds selected from the group consisting of pyrethroids, pyrethroid-type compounds, recombinant nucleopolyhedroviruses capable of expressing an insect toxin, organophosphates, carbamates, formamidines, macrocyclic lactones, amidinohydrazones, GABA (gamma-aminobutyric acid) antagonists, and acetylcholine receptor ligands.

The present invention also provides a method for synergistic insect control which comprises contacting said insect with a synergistically effective amount of a neuronal sodium channel antagonist in combination with one or more compounds selected from the group consisting of pyrethroids, pyrethroid-type compounds, recombinant nucleopolyhedroviruses capable of expressing an insect toxin, organophosphates, carbamates, formamidines, macrocyclic lactones, amidinohydrazones, GABA antagonists and acetylcholine receptor ligands.

The present invention further provides a method for the enhanced protection of plants from infestation and attack by insects.

DETAILED DESCRIPTION OF THE INVENTION

Definitions

" Acetylcholine receptor ligand compound " as used in this application means a compound which is capable of binding to the acetylcholine receptor site.

" Group A " as used in this application means insecticidal

- 1) pyrethroid compounds;
- 2) pyrethroid-type compounds;
- 10 3) recombinant nucleopolyhedroviruses capable of expressing an insect toxin;
- 4) organophosphate compounds;
- 5) carbamate compounds;
- 6) formamidine compounds;
- 15 7) macrocyclic lactone compounds;
- 8) amidinohydrazone compounds;
- 9) GABA antagonist compounds; and
- 10) acetylcholine receptor ligand compounds.

" Haloalkyl" as used in this application means an alkyl group C_xH_{2x+1} having 1 to $2x+1$ halogen atoms which may be the same or different. Similarly, the terms " haloalkenyl" , " haloalkynyl" , " haloalkoxy " , " halophenyl" and the like mean mono- to perhalogen substitution wherein the halogens may be the same or different.

" Halogen " as used in this application means Cl, Br, I or F.

" Neuronal sodium channel antagonist" as used in this application means a compound which is capable of preventing the ability of a neuron cell to transfer sodium ions across the cell membrane.

" Pyrethroid-type compounds " as used in this application means those compounds characterized by a non-ester linked aryl-phenoxybenzyl moiety.

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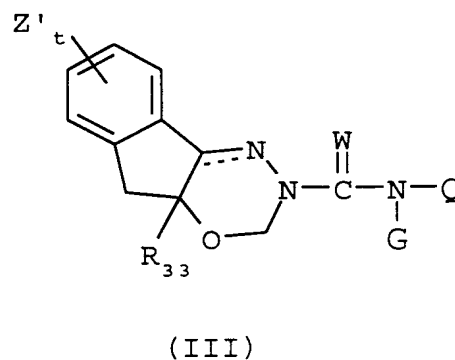
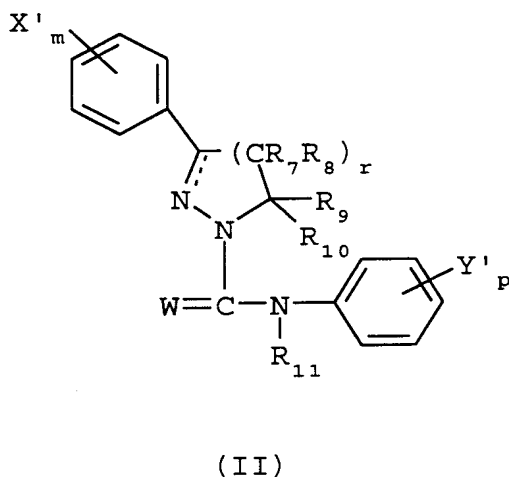
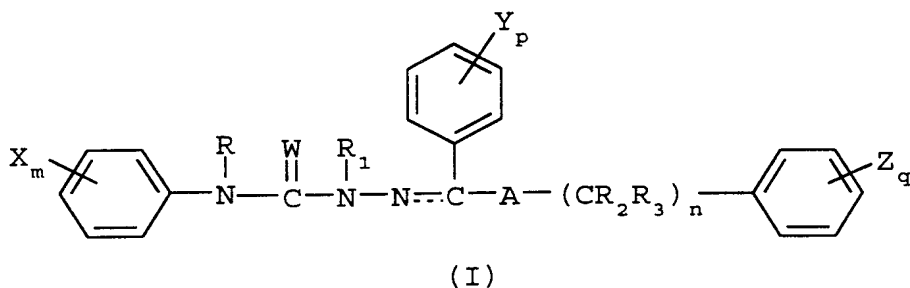
" Synergism " as used in this application means a cooperative action encountered in a combination of two or more biologically active components in which the combined activity of the two or more components exceeds
5 the sum of the activity of each component alone.

Surprisingly, it has now been found that a composition which comprises a combination of a neuronal sodium channel antagonist and a second insecticidal ingredient provides superior insect control at lower
10 levels of the combined active agents than may be achieved when the neuronal sodium channel antagonist or the second insecticidal ingredient is applied alone.

As previously stated, the term neuronal sodium channel antagonist designates a compound which is
15 capable of preventing the ability of a neuron cell to transfer sodium ions across the cell membrane. A neuron cell thus affected is unable to fire, resulting in paralysis, and ultimately mortality, in the target host. Descriptions of neuronal sodium channel
20 antagonists and their mode of action may be found in Pesticide Biochemistry and Physiology, 60: 177-185 or Archives of Insect Biochemistry and Physiology, 37: 91-103.

Neuronal sodium channel antagonists include
25 compounds such as those described in U.S. 5,543,573; U.S. 5,708,170; U.S. 5,324,837 and U.S. 5,462,938, among other publications. Exemplary of the neuronal sodium channel antagonist compounds useful in the composition of this invention are those compounds
30 having the structural formula

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wherein A is CR_4R_5 or NR_6 ;

5 W is O or S;

 X, Y, Z, X', Y' and Z' are each independently H;
 halogen; OH; CN; NO_2 ; C_1 - C_6 alkyl optionally
 substituted with one or more halogen, C_1 -
 C_3 alkoxy, C_1 - C_3 haloalkoxy, C_3 -
 10 C_6 cycloalkyl, C_2 - C_6 alkenyloxy or
 sulfonyloxy groups;
 C_1 - C_6 alkoxy optionally substituted with one
 or more halogen, C_1 - C_3 alkoxy or C_3 -
 C_6 cycloalkyl groups;
 15 C_1 - C_6 alkoxycarbonyl, C_3 -
 C_6 cycloalkylcarbonyloxy, phenyl optionally
 substituted with one or
 more halogen, C_1 - C_4 alkyl, or C_1 - C_4 alkoxy
 groups;

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aminocarbonyloxy optionally substituted with
one or more C₁-C₃alkyl groups;
C₁-C₆alkoxycarbonyloxy; C₁-C₆alkylsulfonyloxy;
C₂-C₆alkenyl; or NR₁₂R₁₃;

5 m, p and q are each independently an integer of 1,
2, 3, 4, or 5;
n is an integer of 0, 1 or 2;
r is an integer of 1 or 2;
t is an integer of 1, 2, 3 or 4;

10 R, R₁, R₂, R₃, R₄ and R₅ are each independently H or
C₁-C₄alkyl;
R₆ is H, C₁-C₆alkyl, C₁-C₆haloalkyl, C₁-C₆alkoxyalkyl,
C₁-C₆alkoxy, C₁-C₆haloalkoxy, C₂-C₆alkenyl,
C₂-C₆alkynyl, C₁-C₆alkylcarbonyl, C₁-C₆alkoxy-
15 carbonyl, C₁-C₆alkylthio, or C₁-
C₆haloalkylthio;
R₇ and R₈ are each independently H; halogen;
C₁-C₆alkyl; C₁-C₆alkylcarbonyloxy; or phenyl
optionally substituted with one or more
20 halogen, CN, NO₂, C₁-C₆alkyl, C₂-C₆halo-
alkyl, C₁-C₆alkoxy or C₁-C₆haloalkoxy
groups;
R₉ and R₁₀ are each independently H, or C₁-C₄alkyl;
R₁₁ is H, C₁-C₆alkyl, C₁-C₆haloalkyl, C₁-C₄alkyl-
25 carbonyl, C₁-C₆alkoxycarbonyl, or C₁-C₆halo-
alkoxycarbonyl;
R₁₂ and R₁₃ are each independently H or C₁-C₆alkyl;
G is H; C₁-C₆alkyl optionally substituted with one
or more halogen, C₁-C₄alkoxy, C₁-
30 C₆haloalkoxy, CN, NO₂S(O)_uR₁₄, COR₁₅,
CO₂R₁₆, phenyl or
C₃-C₆cycloalkyl groups;
C₁-C₆alkoxy; C₁-C₆haloalkoxy; CN; NO₂; S(O)_uR₁₇;
COR₁₈; CO₂R₁₉; phenyl optionally substituted
35 with one or more halogen, CN, C₁-C₃halo-

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alkyl, or C₁-C₃haloalkoxy groups;
C₃-C₆cycloalkyl; or phenylthio;
Q is phenyl optionally substituted with one or
more halogen, CN, SCN, NO₂, S(O)_uR₂₀, C1-
5 C₄alkyl,
C₁-C₄haloalkyl, C₁-C₄alkoxyalkyl, C1-C₆alkoxy,
C₁-C₆haloalkoxy, or NR₂₁R₂₂ groups;
u is an integer of 0, 1 or 2;
R₁₄, R₁₅, R₁₆, R₁₈, R₁₉, R₂₁ and R₂₂ are each
10 independently H or C₁-C₆alkyl;
R₁₇ and R₂₀ are each independently C₁-C₆alkyl or
C₁-C₆haloalkyl;
R₃₃ is CO₂R₃₄;
R₃₄ is H, C₁-C₆alkyl, C₁-C₆haloalkyl, phenyl or
15 halophenyl; and the dotted line configuration
C \cdots N represents a double bond or a single bond
(i.e. C-N or C=N); or
a stereoisomer thereof.

Preferred neuronal sodium channel antagonists
20 suitable for use in the composition of the invention
are those compounds of formula I, II or III wherein the
dotted line configuration C \cdots N represents a double
bond.

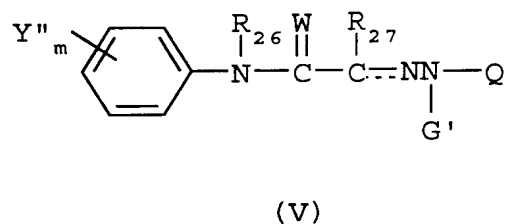
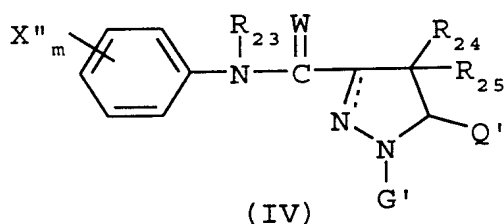
More preferred neuronal sodium channel antagonists
25 suitable for use in the inventive composition are those
compounds of formula I or formula III wherein the
dotted line configuration represents a double bond.

Particularly preferred neuronal sodium channel
antagonists useful in the composition of the invention
30 are those compounds of formula I or formula III wherein
W is O; X is trifluoromethoxy and is in the 4-position;
Y is trifluoromethyl and is in the 3-position; Z is CN
and is in the 4-position; A is CH₂; n is 0; m, p and q
are each 1; R and R₁ are each H; Z₁ is C₁; R₃₃ and G are
35 each CO₂CH₃; Q is p-(trifluoromethoxy)phenyl; and the

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dotted line configuration $C=N$ represents a double bond; or a stereoisomer thereof.

Further neuronal sodium channel antagonist compounds include those described in U.S. 5,116,850 and
 5 U.S. 5,304,573, among other publications. Exemplary of further neuronal sodium channel antagonist compounds suitable for use in the composition of the invention are those compounds having structural formula



10

wherein W is O or S;

X' and Y' are each independently H; halogen; CN;
 15 SCN; C_1 - C_6 alkyl optionally substituted with one or

more halogen, NO_2 , CN, C_1 - C_4 alkoxy,
 C_1 - C_4 alkylthio, phenyl, halophenyl,
 C_1 - C_4 alkylsulfonyl, C_1 -
 20 C_4 haloalkylsulfonyl, or C_1 -
 C_4 alkoxycarbonyl groups;
 C_2 - C_4 alkenyl; C_2 - C_4 haloalkenyl; C_2 - C_4 alkynyl;
 C_2 - C_4 haloalkynyl; C_3 - C_6 cycloalkyl; C_3 -
 C_6 halocycloalkyl; phenyl optionally
 25 substituted
 with one or more halogen, CN, NO_2 , C_1 -
 C_4 alkyl, C_1 - C_4 haloalkyl, C_1 - C_4 alkoxy, C_1 -
 C_4 haloalkoxy, C_1 - C_4 alkylthio, C_1 -
 C_4 alkylsulfonyl or C_1 - C_4 haloalkylsulfonyl
 30 groups;
 C_1 - C_4 alkylcarbonyl; C_1 - C_4 haloalkylcarbonyl; or

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 $NR_{28}R_{29}$;

m is an integer of 1, 2, 3, 4 or 5;

G' is phenyl optionally substituted with one or more groups which may be the same or different selected from X' ;

5

a 5-membered heteroaromatic ring containing one or two heteroatoms selected from 0 or 1 oxygen, 0 or 1 sulfur and 0, 1 or 2 nitrogen atoms said 5-membered heteroaromatic ring being attached via carbon and being optionally substituted with one or more groups which may be the same or different selected from X' ; or

10

a 6-membered heteroaromatic ring containing one or two heteroatoms selected from 0 or 1 oxygen, 0 or 1 sulfur and 0, 1 or 2 nitrogen atoms said 6-membered heteroaromatic ring being attached via carbon and being optionally substituted with one or more groups which may be the same or different selected from X' ;

15

20

Q' is H; C₁-C₆alkyl optionally substituted with one or more halogen, CN, C₁-C₃alkoxy, C₁-C₆alkoxycarbonyl, or phenyl optionally substituted with one or more halogen, CN, NO₂, C₁-C₄alkyl, C₁-C₄haloalkyl, C₁-C₄alkylsulfonyl or C₁-C₄alkylsulfinyl groups;

25

30

C₂-C₆alkenyl; C₂-C₆alkynyl; or phenyl optionally substituted with one to three groups, which may be the same or different, selected from X' ;

R₂₃, R₂₄, R₂₅, R₂₆, R₂₇, R₂₈ and R₂₉ are each

35

independently H or C₁-C₄alkyl; and the dotted

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line configuration $C \equiv N$ represents a double bond or a single bond (i.e. C-N or C=N); or a stereoisomer thereof.

Further preferred neuronal sodium channel
5 antagonist compounds of the invention are those compounds of formula IV or V wherein the dotted line configuration $C \equiv N$ represents a double bond.

Other preferred neuronal sodium channel antagonist
10 compounds suitable for use in the composition of the invention are those compounds of formula IV or V wherein W is O; X' and Y' are each independently H or C₁-C₆haloalkyl; m is 1; R₂₃, R₂₄, R₂₅, R₂₆ and R₂₇ are each H; G is phenyl optionally substituted with one or more halogen atoms; Q' is halophenyl or C₁-C₄alkyl optionally
15 substituted with one phenyl or halophenyl group; and the dotted line configuration $C \equiv N$ represents a double bond; or a stereoisomer thereof.

The second active ingredient of the insecticidal composition of the invention includes one or more
20 compounds selected from Group A:

- 1) pyrethroid compounds which are known to be insecticidally active such as cypermethrin, cyhalothrin, cyfluthrin, permethrin or the like;
- 2) pyrethroid-type compounds which are known to
25 be insecticidally active such as ethofenprox, silafluofen, or the like;
- 3) recombinant nucleopolyhedroviruses capable of expressing an insect toxin, preferably an insect neurotoxin such as Androctonus australis insect toxin
30 (AaIT), for example HzNPV-AaIT;
- 4) organophosphate compounds which are known to be insecticidally active such as profenofos, acephate, sulprofos, malathion, diazinon, methyl parathion, terbufos, or the like;

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5) carbamate compounds which are known to be insecticidally active such as methomyl, thiodicarb, fenothiocarb, or the like;

6) formamidine compounds which are known to be insecticidally active such as amitraz, chlordimeform, hydramethylnon, chlorfenamidin, or the like;

7) macrocyclic lactone compounds which are known to be insecticidally active such as spinosad, avermectin, emamectin, milbemectin, nemadectin, moxidectin or the like;

8) amidinohydrazone compounds which are known to be insecticidally active such as hydramethylnon;

9) GABA antagonist compounds which are known to be insecticidally effective such as fipronil, endosulfan, or the like;

10) acetylcholine receptor ligand compounds which are known to be insecticidally effective such as imidacloprid, acetamiprid, nitenpyram, thiamethoxam, or the like.

20 Descriptions of the above-listed commercially available compounds may be found in The Pesticide Manual, 11th Edition, British Crop Protection Council (1997) among other publications. Descriptions of recombinant nucleopolyhedroviruses capable of expressing an insect toxin include Treacy *et al*,
25 Proceedings Beltwide Cotton Conference (1999), pp 1076-1083.

Preferred compositions of the invention are those compositions having a neuronal sodium channel antagonist compound of formula I or formula III in combination with one or more compounds selected from Group A.

More preferred compositions of the invention are those compositions having a formula I or formula III compound wherein W is O; X is trifluoromethoxy and is in
35

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the 4-position; Y is trifluoromethyl and is in the 3-position; Z is CN and is in the 4-position; A is CH₃; n is 0; m, p and q are each independently 1; R and R₁ are each independently H; Z' is Cl; R₃₃ and G are each independently CO₂CH₃; Q is p-(trifluoromethoxy)phenyl; and the dotted line configuration C=N represents a double bond in combination with one or more compounds selected from Group A.

Each of the compounds of formula I, II, III, IV and V embody assymetric centers which may be represented in the stereoisomeric R-form or S-form. The present invention also includes the R-form, the S-form or mixtures comprising the R-form and the S-form in an arbitrary ratio. For compounds of formula III, the S-form is preferred.

Advantageously, the neuronal sodium-channel antagonist compound of formula I, II, III, IV or V or a mixture thereof may be formulated with a second insecticidally effective ingredient and optionally other customary formulation adjuvants. Said formulation may be dispersed in a solid or liquid diluent for application to the insect, its food supply, breeding ground or habitat as a dilute spray or as a solid dust or dust concentrate.

The active ingredients of the inventive composition may also be formulated separately as a wettable powder, emulsifiable concentrate, aqueous or liquid flowable, suspension concentrate or any one of the conventional formulations used for insect control agents and tank-mixed in the field with water or other inexpensive liquid for application as a liquid spray mixture. The separately formulated compositions may also be applied sequentially.

Advantageously, the composition of the invention may be formulated as a bait composition comprising a

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synergistically effective amount of a combination of a neuronal sodium channel antagonist plus one or more compounds selected from Group A and a solid or liquid edible nutritive substance. A preferred bait
5 composition may contain by weight about 0.01% to 20% active ingredients, preferably a neuronal sodium channel antagonist in combination with hydramethylnon.

In actual practice, the composition of the invention may be applied to the plant foliage or plant
10 stem or to the insect habitat or to the locus of a hygienic pest as a dilute spray prepared from any of the above-said formulations. The ratio of the essential active ingredients of the composition of the invention is about 1 weight part of a neuronal sodium
15 channel antagonist to about 0.01-100 weight parts of one or more compounds selected from Group A.

The compositions of the invention are superior insecticidal compositions and are especially useful for the control of agrohorticultural pests, hygienic pests
20 or wood-eating pests. Said compositions are highly effective for the protection of growing and harvested plants including: leguminous crops such as soybeans, snap beans, peas, wax beans and the like as well as cotton, forage crops, cole crops, leafy vegetables,
25 tobacco, hops, tomatoes, potatoes, flowering ornamentals such as chrysanthemums, vine crops such as grapes, squash, pumpkin or melon and fruit trees such as cherry, peach, apple or citrus, from the ravages of insects.

30 The synergistic insecticidal composition of the invention is found to be highly active against a wide variety of lepidopteran and coleopteran insects such as *Helicoverpa zea* (cotton bollworm), *Heliothis virescens* (tobacco budworm), *Leptinotarsa decemlineata* (Colorado

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potato beetle), *Diabrotica spp.* (corn rootworm) and the like.

Beneficially, the composition of the invention may be useful for the prevention and control of hygienic or
5 public health pests such as: Diptera, e.g. houseflies, mosquitoes, or the like; Hymenoptera, e.g. ants, parasitic wasps, wasps or the like; Blattaria, e.g. cockroaches; or the like.

Further, the compositions of the invention may be
10 particularly useful for the prevention and control of wood-eating insects such as termites (Isoptera), carpenter ants (Hymenoptera), wood-destroying beetles (Coleoptera) or the like.

These and other advantages of the invention may
15 become more apparent from the examples set forth herein below. These examples are provided merely as illustrations of the invention and are not intended to be construed as a limitation thereof.

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EXAMPLE 1Evaluation of the Synergistic Insecticidal Effect
of a Combination of a Neuronal Sodium Channel5 Antagonist Plus a Second Insecticide

In this evaluation, *Heliothis zea* (cotton bollworm), *Heliothis virescens* (tobacco budworm) and pyrethroid-resistant *Heliothis virescens* larvae used are obtained from laboratory colonies. Pyrethroid-resistant
10 *H. virescens* are derived from the PEG-strain [Campannola & Plapp, Proceedings of Beltwide Cotton Conference (1988)].

Cotton leaves are immersed in 1:1 v/v, acetone/water solutions of test compound, or solutions
15 of a combination of test compounds for a period of about 3 seconds. Following immersion, leaves are allowed to air-dry for 2-3 hours. Plastic bioassay trays containing multiple open-faced wells (4.0 x 4.0 x 2.5
cm) are used as the test arenas. Cut portions of a
20 treated leaf, a moistened cotton dental wick and a single third-instar larva are placed into each well, covered with an adhesive vented clear plastic sheet and held under constant fluorescent light at about 27°C for a predetermined period of time. Larval
25 mortality/morbidity is evaluated at 5 days after treatment. All treatments are replicated 4-5 fold in a randomized complete block design with 16-32 larvae per treatment. Using conventional log-probit analysis, the LC₅₀ of each treatment is determined.

30 Using the above protocol, a neuronal sodium channel antagonist (Compound A) may be evaluated alone at dose rates of 0.1 ppm, 1.0 ppm and 10.0 ppm and in combination with 1.0 ppm of a second insecticidal

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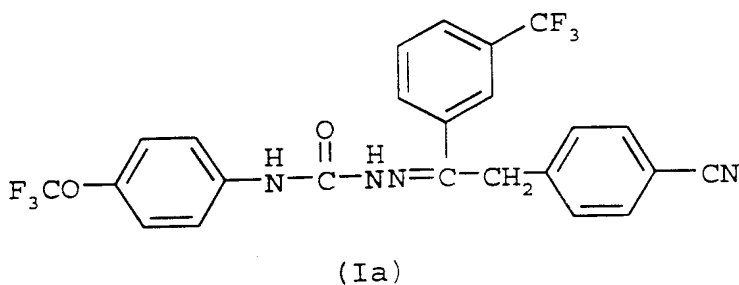
compound. Treatments which may be used are shown in Table I.

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Table I

Second Active Compound	Dose Rate (ppm)	Compound A1			
		Dose Rate			
		(ppm)	(ppm)	(ppm)	(ppm)
cypermethrin	0	0	0.1	1.0	10.0
	1.0	0	0.1	1.0	10.0
amitraz	0	0	0.1	1.0	10.0
	1.0	0	0.1	1.0	10.0
fipronil	0	0	0.1	1.0	10.0
	1.0	0	0.1	1.0	10.0
acetamiprid	0	0	0.1	1.0	10.0
	1.0	0	0.1	1.0	10.0
spinosad	0	0	0.1	1.0	10.0
	1.0	0	0.1	1.0	10.0
thiodicarb	0	0	0.1	1.0	10.0
	1.0	0	0.1	1.0	10.0

¹Compound A = formula Ia neuronal sodium channel antagonist



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EXAMPLE 2

Evaluation of the Synergistic Insecticidal Effect Of a
Combination Of A Neuronal Sodium Channel Antagonist
5 Plus an Amidinohydrazone

In this evaluation, adult male German cockroaches (*Blattella germanica*) are used. For each test, a 4.0 g portion of ground Purina Dog Chow (Hi-Pro Glo[®]) is treated with an acetone solution of test compound alone
10 or in combination with a second test compound. After treatment, the acetone is evaporated and the treated dog chow is placed in a 3/4 oz plastic cup which is placed in a harborage made of folded sheets of blotter paper placed in a plastic box (16" L x 11" W x 6" H).
15 The plastic box (test arena) is also fitted with a 1 oz narrow mouth bottle with 2 dental wicks inserted at the mouth. A control box is prepared in the same manner using ground dog chow which has been treated with reagent grade acetone. Each treatment is replicated
20 three times. Into each test arena are placed 20 healthy adult male cockroaches which have been reared in an insectary. The test arenas are then stored at 76°F and mortality is determined daily by visual examination. The data obtained are shown in Table II.

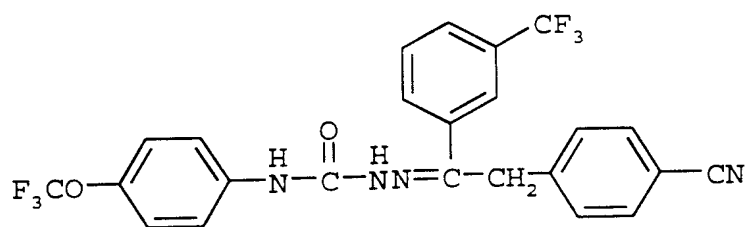
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Table II

Test	% Active	% Mortality					
		Days After Treatment					
<u>Compound</u>	<u>Ingredien</u>	<u>3</u>	<u>4</u>	<u>5</u>	<u>6</u>	<u>7</u>	<u>8</u>
A ¹	0.05	0	0	0	0	0	0
A	0.10	1.7	11.7	11.7	11.7	18.3	18.3
A	0.50	5.0	5.0	5.0	5.0	5.0	5.0
B ²	1.00	0	5.0	28.3	71.7	90.0	93.3
A+B	0.05+1.0	0	20.0	41.7	81.7	95.0	98.3
A+B	0.10+1.0	0	21.7	51.7	88.3	95.0	95.0
A+B	0.50+1.0	16.7	58.3	80.0	95.0	98.3	100.0
Control	0	0	1.7	3.3	3.3	3.3	5.0

¹Compound A = formula Ia neuronal sodium channel antagonist

²Compound B = hydramethylnon



(Ia)

As can be seen from the data shown in Table II, combinations of a neuronal sodium channel antagonist plus an amidinohydrazone insecticide demonstrate synergistic insect control.

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EXAMPLE 3Evaluation of the Synergistic Insecticidal Effect Of a Combination Of A Neuronal Sodium Channel Antagonist Plus A Recombinant Nucleopolyhedrovirus Capable Of Expressing An Insect Toxin

In this evaluation, *Helicoverpa zea* (cotton bollworm) larvae are obtained from a laboratory colony. Test compounds are dissolved in 1:1 v/v acetone/water. Plastic bioassay trays (C-D International, Pitman, NJ) are used as test arenas. Each tray contains 32 open-faced wells, 4.0 x 4.0 x 2.5 cm. A portion (5 ml) of a wheat germ-soybean flour-based artificial diet (Southland Products, Lake Village, AR) is poured into each well. After the diet hardened, 0.4 ml of test solution is pipetted onto the diet surface in each well. Test solutions are evenly spread over surfaces of diet by picking up the tray and gently tilting it from side to side. Trays are then held in a vented area for about 2 h, until water is no longer pooled on diet surfaces. A single 4-day-old *H. zea* larva is then placed on the surface of diet in each well. After larval infestation, each well is covered with an adhesive, vented, clear plastic sheet.

All test arenas are held under constant fluorescent light and a temperature of about 27°C for duration of the assay. Larval mortality is determined at 2, 3, 4 and 7 days after treatment. A larva was considered to be dead if it exhibited little to no movement, even after being shaken in the diet tray. A total of 32 insects were tested for each treatment.

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The data obtained are shown in Table III.

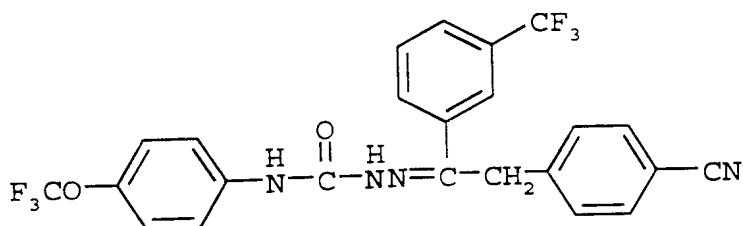
Table III

Test Compound	Conc. of Active Ingredient	% Mortality			
		Days After Treatment			
		2	3	4	7
A ¹	0.1 ppm	43.8	46.9	53.1	53.1
B ²	1000 OB ³ /ml	3.1	34.4	50.0	62.5
B	500 OB/ml	0.0	9.4	18.8	40.6
B	100 OB/ml	3.1	3.1	3.1	15.6
A+B	0.1+1000	87.5	90.6	93.8	96.9
A+B	0.1+500	75.0	78.1	84.4	87.5
A+B	0.1+100	62.5	75.0	75.0	78.1
Control	0	3.1	3.1	3.1	3.1

¹Compound A = formula Ia neuronal sodium channel antagonist

²Compound B = HzNPV-AaIT, *Helicoverpa zea* Nucleopolyhedrovirus
which expresses *Androctonus australis* insect toxin

³OB = viral occlusion bodies



(Ia)

5

As can be seen from the data shown in Table III, combinations of a neuronal sodium channel antagonist plus a recombinant nucleopolyhedrovirus which is

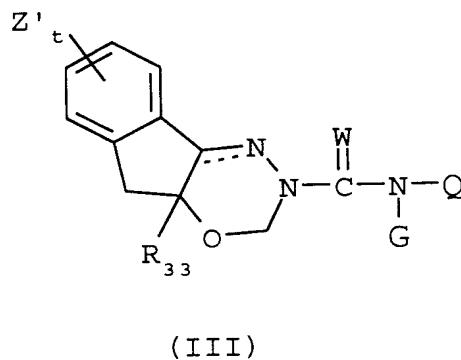
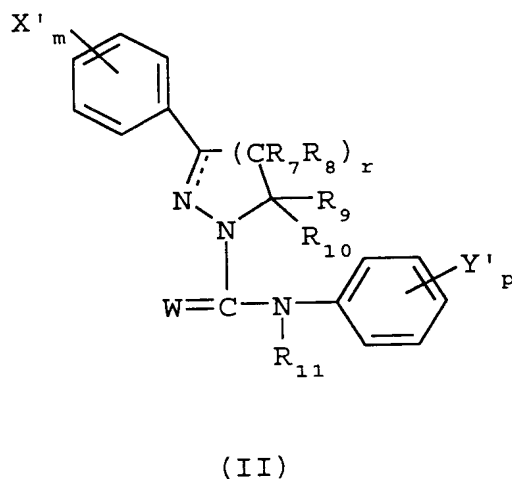
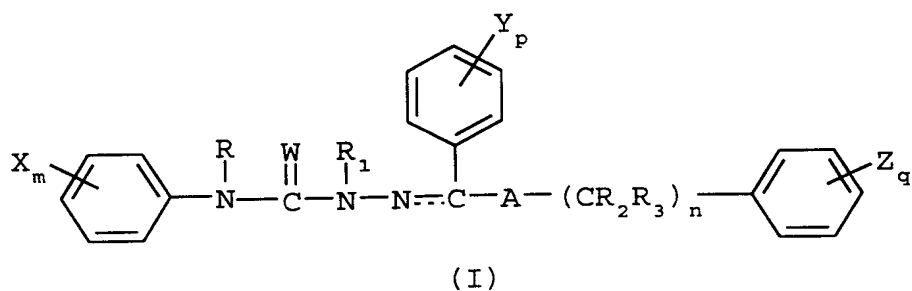
-22-

capable of expressing an insect toxin demonstrate synergistic insect control.

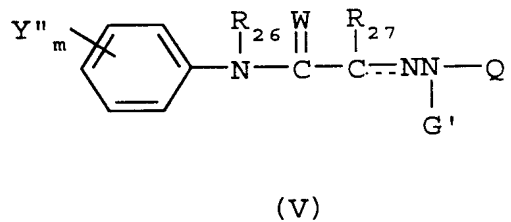
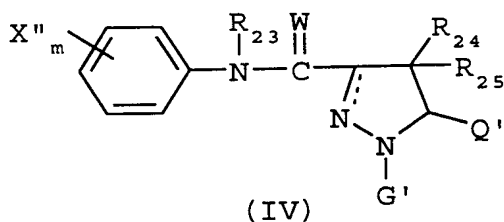
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WHAT IS CLAIMED IS:

1. A synergistic insecticidal composition comprising a synergistically effective amount of a neuronal sodium channel antagonist in combination with one or more compounds selected from Group A wherein the neuronal sodium channel antagonist is a compound of formula



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wherein A is CR_4R_5 or NR_6 ;

W is O or S;

X, Y, Z, X', Y' and Z' are each independently H; halogen; OH; CN; NO_2 ; C_1 - C_6 alkyl optionally substituted with one or more halogen,

C_1 - C_3 alkoxy, C_1 - C_3 haloalkoxy, C_3 - C_6 cycloalkyl, C_2 - C_6 alkenyloxy or sulfonyloxy groups;

C_1 - C_6 alkoxy optionally substituted with one or more halogen, C_1 - C_3 alkoxy or C_3 - C_6 cycloalkyl groups;

C_1 - C_6 alkoxycarbonyl, C_3 - C_6 cycloalkylcarbonyloxy, phenyl optionally substituted with one or more halogen, C_1 - C_4 alkyl, or C_1 - C_4 alkoxy groups;

aminocarbonyloxy optionally substituted with one or more C_1 - C_3 alkyl groups;

C_1 - C_6 alkoxycarbonyloxy; C_1 - C_6 alkylsulfonyloxy; C_2 - C_6 alkenyl; or $NR_{12}R_{13}$;

m, p and q are each independently an integer of 1, 2, 3, 4, or 5;

n is an integer of 0, 1 or 2;

r is an integer of 1 or 2;

t is an integer of 1, 2, 3 or 4;

R , R_1 , R_2 , R_3 , R_4 and R_5 are each independently H or C_1 - C_4 alkyl;

R_6 is H, C_1 - C_6 alkyl, C_1 - C_6 haloalkyl, C_1 - C_6 alkoxyalkyl, C_1 - C_6 alkoxy, C_1 - C_6 haloalkoxy, C_2 - C_6 alkenyl,

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C_2-C_6 alkynyl, C_1-C_6 alkylcarbonyl, C_1-C_6 alkoxy-carbonyl, C_1-C_6 alkylthio, or C_1-C_6 haloalkylthio;

R_7 and R_8 are each independently H; halogen; C_1-C_6 alkyl; C_1-C_6 alkylcarbonyloxy; or phenyl optionally substituted with one or more halogen, CN, NO_2 , C_1-C_6 alkyl, C_2-C_6 haloalkyl, C_1-C_6 alkoxy or C_1-C_6 haloalkoxy groups;

R_9 and R_{10} are each independently H, or C_1-C_4 alkyl; R_{11} is H, C_1-C_6 alkyl, C_1-C_6 haloalkyl, C_1-C_4 alkylcarbonyl, C_1-C_6 alkoxycarbonyl, or C_1-C_6 haloalkoxycarbonyl;

R_{12} and R_{13} are each independently H or C_1-C_6 alkyl; G is H; C_1-C_6 alkyl optionally substituted with one or more halogen, C_1-C_4 alkoxy, C_1-C_6 haloalkoxy, CN, $NO_2S(O)_uR_{14}$, COR_{15} , CO_2R_{16} , phenyl or C_3-C_6 cycloalkyl groups;

C_1-C_6 alkoxy; C_1-C_6 haloalkoxy; CN; NO_2 ; $S(O)_uR_{17}$; COR_{18} ; CO_2R_{19} ; phenyl optionally substituted with one or more halogen, CN, C_1-C_3 haloalkyl, or C_1-C_3 haloalkoxy groups; C_3-C_6 cycloalkyl; or phenylthio;

Q is phenyl optionally substituted with one or more halogen, CN, SCN, NO_2 , $S(O)_uR_{20}$, C_1-C_4 alkyl, C_1-C_4 haloalkyl, C_1-C_4 alkoxyalkyl, C_1-C_6 alkoxy, C_1-C_6 haloalkoxy, or $NR_{21}R_{22}$ groups;

u is an integer of 0, 1 or 2;

R_{14} , R_{15} , R_{16} , R_{18} , R_{19} , R_{21} and R_{22} are each independently H or C_1-C_6 alkyl;

R_{17} and R_{20} are each independently C_1-C_6 alkyl or C_1-C_6 haloalkyl;

R_{33} is CO_2R_{34} ;

R_{34} is H, C_1-C_6 alkyl, C_1-C_6 haloalkyl, phenyl or

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halophenyl;

X' and Y' are each independently H; halogen; CN; SCN; C₁-C₆alkyl optionally substituted with one

or more halogen, NO₂, CN, C₁-C₄alkoxy, C₁-C₄alkylthio, phenyl, halophenyl, C₁-C₄alkylsulfonyl, C₁-C₄haloalkylsulfonyl, or C₁-C₄alkoxycarbonyl groups;

C₂-C₄alkenyl; C₂-C₄haloalkenyl; C₂-C₄alkynyl; C₂-C₄haloalkynyl; C₃-C₆cycloalkyl; C₃-C₆halo-cycloalkyl; phenyl optionally substituted

with one or more halogen, CN, NO₂, C₁-C₄alkyl, C₁-C₄haloalkyl, C₁-C₄alkoxy, C₁-C₄haloalkoxy, C₁-C₄alkylthio, C₁-C₄alkylsulfonyl or

C₁-C₄haloalkylsulfonyl groups;

C₁-C₄alkylcarbonyl; C₁-C₄haloalkylcarbonyl; or NR₂₈R₂₉;

G' is phenyl optionally substituted with one or more groups which may be the same or different selected from X' ;

- a 5-membered heteroaromatic ring containing one or two heteroatoms selected from 0 or 1 oxygen, 0 or 1 sulfur and 0, 1 or 2 nitrogen atoms said 5-membered heteroaromatic ring being attached via carbon and being optionally substituted with one or more groups which may be the same or different selected from X' ; or
- a 6-membered heteroaromatic ring containing one or two heteroatoms selected from 0 or 1 oxygen, 0 or 1 sulfur and 0, 1 or 2 nitrogen atoms said 6-membered heteroaromatic ring being attached via carbon and being optionally substituted

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with one or more groups which may be the same or different selected from X' ;

Q' is H; C₁-C₆alkyl optionally substituted with one or more halogen, CN, C₁-C₃alkoxy, C₁-C₆alkoxycarbonyl, or phenyl optionally

substituted with one or more halogen, CN, NO₂, C₁-C₄alkyl, C₁-C₄haloalkyl, C₁-C₄alkylsulfonyl or C₁-C₄alkylsulfinyl groups;

C₂-C₆alkenyl; C₂-C₆alkynyl; or phenyl optionally substituted with one to three groups, which may be the same or different, selected from X' ;

R₂₃, R₂₄, R₂₅, R₂₆, R₂₇, R₂₈ and R₂₉ are each independently H or C₁-C₄alkyl; and the dotted line configuration C=N represents a double bond or a single bond; or a stereoisomer thereof.

2. The composition according to claim 1 wherein the neuronal sodium channel antagonist is a compound of formula I or III and the dotted line configuration C=N represents a double bond.

3. The composition according to claim 2 wherein W is O; X is trifluoromethoxy and is in the 4-position; Y is trifluoromethyl and is in the 3-position; Z is CN and is in the 4-position; A is CH₂; n is 0; m, p and q are each 1; R and R₁ are each H; Z' is Cl; R₃₃ and G are each CO₂CH₃; and Q is p-(trifluoromethoxy)phenyl.

4. The composition according to claim 3 wherein the one or more compounds selected from Group A are

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cypermethrin, cyhalomethrin, cyfluthrin, permethrin, ethofenprox, silafluofen, fipronil, endosulfon, imidacloprid, acetamiprid, nitenpyram, thiamethoxam, profenofos, acephate, sulprofos, malathion, diazinon, methyl parathion, terbufos, methonyl, thiodicarb, fenothiocarb, amitraz, chlordimeform, chlorfenamidin, avermectin, emamectin, milbemectin, nemadectin, or moxidectin.

5. The composition according to claim 3 wherein the one or more compounds selected from Group A is a recombinant nucleopolyhedrovirus capable of expressing insect toxin.

6. The composition according to claim 3 wherein the one or more compounds selected from Group A is hydramethylnon.

7. A method for synergistic insect control which comprises contacting said insect with a composition of any one of claims 1-6.

8. The method according to claim 7 wherein the insect is selected from the group consisting of Blattaria, Isoptera, Diptera, and Hymenoptera.

9. The method according to claim 8 wherein the insects are lepidoptera or coleoptera.

10. A method for protecting a plant from infestation and attack by insects which comprises applying to the foliage or stem of said plant a synergistically effective amount of a composition according to any one of claims 1-6.